

AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application:

List of Claims:

Claims 1- 27 (cancelled)

Claim 28 (Currently amended): A method of simultaneously genotyping multiple samples in a single round of hybridization, the method comprising:

- 1) incubating a microarray of polynucleotide samples from multiple individuals with a single solution of a probe mixture of oligonucleotides of known sequence, wherein
 - a) the microarray contains a plurality of samples containing genotypes of interest with each sample in a distinct location ~~and, each location~~ occupying an area smaller than or about 1 square millimeter across,
 - b) each sample has amplified polynucleotides with a defined segment containing a marker selected from a marker for a gene and markers for allelic variants of the gene,
 - c) the oligonucleotides in the probe mixture are of known sequence and length and have sequences specifically complementary to polynucleotide sequences within the defined segments for each sample for which a genotype is to be determined, wherein the oligonucleotides complementary to the polynucleotides are selected from the group consisting of oligonucleotides with sequences complementary to a segment containing the marker for (1) a gene, (2) one or more allelic variants of the gene, and (3) a gene and one or more allelic variants of the gene, and
 - d) the incubating forms hybrids of polynucleotides of the microarray and complementary oligonucleotides and allows discrimination at single nucleotide resolution; and
- 2) detecting at the distinct ~~location~~ locations on the microarray after a single round of hybridization, stable hybrids formed during the incubation, wherein a hybridization signal indicating the formation of a hybrid or lack of formation of a hybrid genotypes the ~~individual~~ individuals.

Claim 29 (previously presented): The method of claim 28 wherein the polynucleotide samples of the microarray are amplification products.

Claim 30 (previously presented): The method of claim 29, wherein the amplification products are produced by a polymerase chain reaction (PCR) method.

Claim 31 (previously amended): The method of claim 30 wherein the plurality of samples of polynucleotides is at least 10.

Claim 32 (previously presented): The method of claim 28 wherein an allele of the gene is associated with a disease.

Claim 33 (previously presented): The method of claim 32 wherein the disease is a human disease.

Claim 34 (previously presented): The method of claim 32 wherein the gene is human and is selected from the group consisting of β -globin, Cystic Fibrosis Transmembrane Conductance Regulator (CFTR), and Galactose-1-Phosphate Uridyltransferase (Gal-1-PU).

Claim 35 (previously presented): The method of claim 28 wherein the microarray is on a surface containing at least 1000 locations per square centimeter.

Claim 36 (previously amended): The method of claim 28 wherein the probe mixture of oligonucleotides of known sequence comprises oligonucleotides with ten different sequences.

Claim 37 (previously presented): The method of claim 28 wherein the oligonucleotides in the mixture are between about 10 and 30 nucleotides in length.

Claim 38 (previously presented): The method of claim 28 wherein the distinct segment is between about 40 and about 1000 nucleotides.

Claim 39 (previously presented): The method of claim 28 wherein the incubating is in an aqueous solution comprised of salts and detergent.

Claim 40 (previously presented): The method of claim 28 wherein hybridizing is performed at a temperature about 10 °C below the melting temperature of the stable hybrids.

Claim 41 (previously presented): The method of claim 28 wherein the oligonucleotides of known sequence are labeled.

Claim 42 (previously presented): The method of claim 41 wherein the label is fluorescent.

Claim 43 (previously presented): The method of claim 28, wherein samples from homozygotes and samples from heterozygotes are distinguishable.

Claim 44 (previously amended): The method of claim 28 wherein the plurality of samples of polynucleotides is at least 5,000.

Claim 45 (previously presented): The method of claim 28 wherein the individual specimens are neonatal blood samples.

Claim 46 (previously amended): The method of claim 28 wherein the individual is a human.

Claim 47 (New): A method of simultaneously genotyping multiple samples in a single round of hybridization, the method comprising:

- 1) incubating a microarray of polynucleotide samples from multiple individuals with a single solution of a probe mixture of oligonucleotides of known sequence, wherein
 - a) the microarray contains a plurality of samples containing genotypes of interest with each sample in a distinct location, wherein the microarray contains at least 60 sample locations per cm^2 ,

- b) each sample has amplified polynucleotides with a defined segment containing a marker selected from a marker for a gene and markers for allelic variants of the gene,
 - c) the oligonucleotides in the probe mixture are of known sequence and length and have sequences specifically complementary to polynucleotide sequences within the defined segments for each sample for which a genotype is to be determined, wherein the oligonucleotides complementary to the polynucleotides are selected from the group consisting of oligonucleotides with sequences complementary to a segment containing the marker for (1) a gene, (2) one or more allelic variants of the gene, and (3) a gene and one or more allelic variants of the gene, and
 - d) the incubating forms hybrids of polynucleotides of the microarray and complementary oligonucleotides and allows discrimination at single nucleotide resolution; and
- 2) detecting at the distinct locations on the microarray after a single round of hybridization, stable hybrids formed during the incubation, wherein a hybridization signal indicating the formation of a hybrid or lack of formation of a hybrid genotypes the individuals.

Claim 48 (New): A method of simultaneously genotyping multiple samples in a single round of hybridization, the method comprising:

- 1) incubating a microarray of polynucleotide samples from multiple individuals with a single solution of a probe mixture of oligonucleotides of known sequence, wherein
- a) the microarray contains a plurality of samples containing genotypes of interest with each sample in a distinct location on an impermeable support,
 - b) each sample has amplified polynucleotides with a defined segment containing a marker selected from a marker for a gene and markers for allelic variants of the gene,
 - c) the oligonucleotides in the probe mixture are of known sequence and length and have sequences specifically complementary to polynucleotide sequences within the defined segments for each sample for which a genotype is to be determined, wherein the oligonucleotides complementary to the polynucleotides are selected from the group consisting of oligonucleotides with sequences complementary to a segment containing the marker for (1) a gene, (2) one or more allelic variants of the gene, and (3) a gene and one or more allelic variants of the gene, and

- d) the incubating forms hybrids of polynucleotides of the microarray and complementary oligonucleotides and allows discrimination at single nucleotide resolution; and
- 2) detecting at the distinct locations on the microarray after a single round of hybridization, stable hybrids formed during the incubation, wherein a hybridization signal indicating the formation of a hybrid or lack of formation of a hybrid genotypes the individuals.

Claim 49 (New): The method of claim 48, wherein the impermeable support further comprises an impermeable surface.

Claim 50 (New): The method of claim 48, wherein the impermeable support further comprises a permeable surface.

Claim 51 (New): The method of claim 48, wherein the impermeable support is rigid.

Claim 52 (New): The method of claim 48, wherein the impermeable support further comprises a surface comprising a reactive group that allows specific attachment of the amplified polynucleotides to the support.

Claim 53 (New): The method of claim 48 wherein the polynucleotide samples of the microarray are amplification products.

Claim 54 (New): The method of claim 53, wherein the amplification products are produced by a polymerase chain reaction (PCR) method.

Claim 55 (New): The method of claim 54 wherein the plurality of samples of polynucleotides is at least 10.

Claim 56 (New): The method of claim 48 wherein an allele of the gene is associated with a disease.

Claim 57 (New): The method of claim 56 wherein the gene is human and is selected from the group consisting of β -globin, Cystic Fibrosis Transmembrane Conductance Regulator (CFTR), and Galactose-1-Phosphate Uridyltransferase (Gal-1-PU).

Claim 58 (New): The method of claim 48 wherein the microarray is on a surface containing at least 1000 sample locations per square centimeter.

Claim 59 (New): The method of claim 48 wherein the probe mixture of oligonucleotides of known sequence comprises oligonucleotides with ten different sequences.

Claim 60 (New): The method of claim 48 wherein the oligonucleotides in the mixture are between about 10 and 30 nucleotides in length.

Claim 61 (New): The method of claim 48 wherein the distinct segment is between about 40 and about 1000 nucleotides.

Claim 62 (New): The method of claim 48 wherein hybridizing is performed at a temperature about 10°C below the melting temperature of the stable hybrids.

Claim 63 (New): The method of claim 48 wherein the oligonucleotides of known sequence are labeled.

Claim 64 (New): The method of claim 63 wherein the label is fluorescent.

Claim 65 (New): The method of claim 48, wherein samples from homozygotes and samples from heterozygotes are distinguishable.

Claim 66 (New): The method of claim 48 wherein the plurality of samples of polynucleotides is at least 5,000.

Claim 67 (previously presented): The method of claim 48 wherein the individual specimens are neonatal blood samples.

Claim 68 (New): The method of claim 48 wherein the individual is a human.